

PATENT COOPERATION TREATY

From the
INTERNATIONAL SEARCHING AUTHORITY

To:
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PCT

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

(PCT Rule 43bis.1)

Applicant's or agent's file reference 5853-449WO		Date of mailing (day/month/year) 18 OCT 2005 FOR FURTHER ACTION See paragraph 2 below	
International application No. PCT/US05/10112	International filing date (day/month/year) 25 March 2005 (25.03.2005)	Priority date (day/month/year) 26 March 2004 (26.03.2004)	
International Patent Classification (IPC) or both national classification and IPC IPC(7): A61B 5/055 and US CL: 424/9.3, 9.32			
Applicant UNIVERSITY OF FLORIDA RESEARCH FOUNDATION, INC.			

1. This opinion contains indications relating to the following items:

- ☒ Box No. I Basis of the opinion
- ☐ Box No. II Priority
- ☐ Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability
- ☐ Box No. IV Lack of unity of invention
- ☒ Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- ☐ Box No. VI Certain documents cited
- ☐ Box No. VII Certain defects in the international application
- ☐ Box No. VIII Certain observations on the international application

2. FURTHER ACTION

If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis(b) that written opinions of this International Searching Authority will not be so considered.

If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later.

For further options, see Form PCT/ISA/220.

3. For further details, see notes to Form PCT/ISA/220.

Name and mailing address of the ISA/ US Mail Stop PCT, Attn: ISA/US Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450 Facsimile No. (703) 305-3230	Authorized officer Michael G. Hartley <i>James Ford</i> Telephone No. (703) 308-1235
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**WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY**

International application No.

PCT/US05/10112

Box No. 1 Basis of this opinion

1. With regard to the **language**, this opinion has been established on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.

☐ This opinion has been established on the basis of a translation from the original language into the following language _____, which is the language of a translation furnished for the purposes of international search (under Rules 12.3 and 23.1(b)).

2. With regard to any **nucleotide and/or amino acid sequence** disclosed in the international application and necessary to the claimed invention, this opinion has been established on the basis of:

a. type of material

☐ a sequence listing

☐ table(s) related to the sequence listing

b. format of material

☐ in written format

☐ in computer readable form

c. time of filing/furnishing

☐ contained in international application as filed.

☐ filed together with the international application in computer readable form.

☐ furnished subsequently to this Authority for the purposes of search.

3. ☐ In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished.

4. Additional comments:

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

International application No.
PCT/US05/10112

Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)	Claims <u>NONE</u>	YES
	Claims <u>1-31</u>	NO
Inventive step (IS)	Claims <u>NONE</u>	YES
	Claims <u>1-31</u>	NO
Industrial applicability (IA)	Claims <u>1-31</u>	YES
	Claims <u>NONE</u>	NO

2. Citations and explanations:

Claims 1-31 lack novelty under PCT Article 33(2) as being anticipated by GRINSTAFF (US 5,498,421). GRINSTAFF discloses a method of assessing oxygenation comprising administering a contrast agent comprising PEG or pyridoxal phosphate modified hemoglobin and performing MRI imaging, see column 29, lines 28-42, column 26, lines 212-61 and examples 18-19. The contrast agents may further include various paramagnetic metals to enhance the MRI image, see column 7, lines 11-14. The hemoglobin contrast agents also may include various surfactants as claimed, see column 12, lines 14+. The contrast agents may further include a perfluorocarbon in the core of microparticulate contrast agents, see column 27.

Claims 1-31 lack an inventive step under PCT Article 33(3) as being obvious over GRINSTAFF discloses a method of assessing oxygenation as set forth above. It would have been obvious to one of ordinary skill in the art to employ any of the contrast agents, surfactants, metals, etc., taught by GRINSTAFF because the various components of the contrast agents are taught by GRINSTAFF to be interchangeable for optimizing the imaging capabilities for various purposes, including assessing the oxygenation of tissues.

Claims 1-31 meet the criteria set out in PCT Article 33(4), and thus have industrial applicability because the subject matter claimed can be made or used in industry.